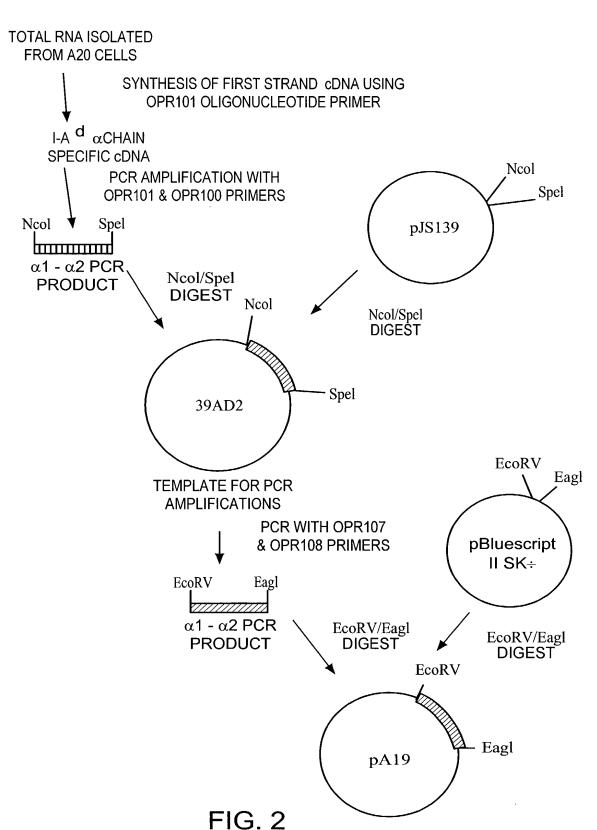


SCHEMATIC VIEW OF SOLUBLE PEPTIDE-LINKED MHC- $\lg G$  C-REGION FUSION PROTEIN

# $^{3/64}$ I-A $^{d}$ $_{\alpha}$ CHAIN CLONING SCHEME



## β CHAIN CLONING SCHEME

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TOTAL RNA ISOLATED FROM A20 CELLS

> SYNTHESIS OF FIRST STRAND cDNA USING OLIGO dT PRIMER

#### cDNA TEMPLATE

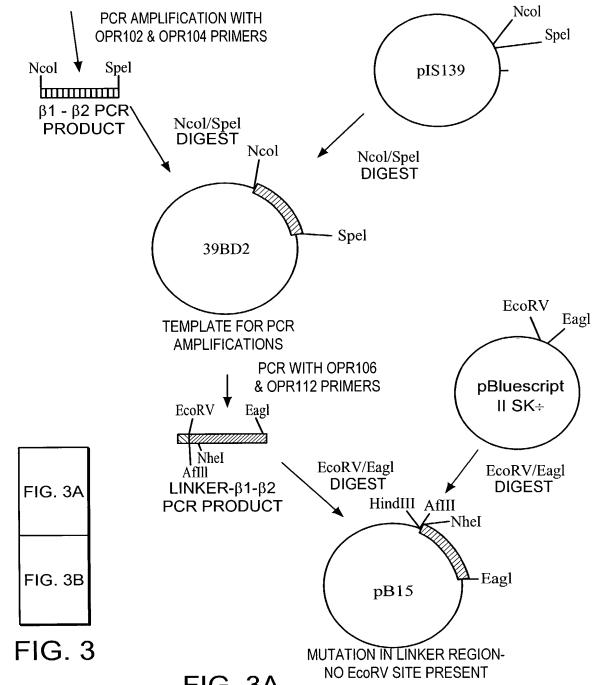


FIG. 3A

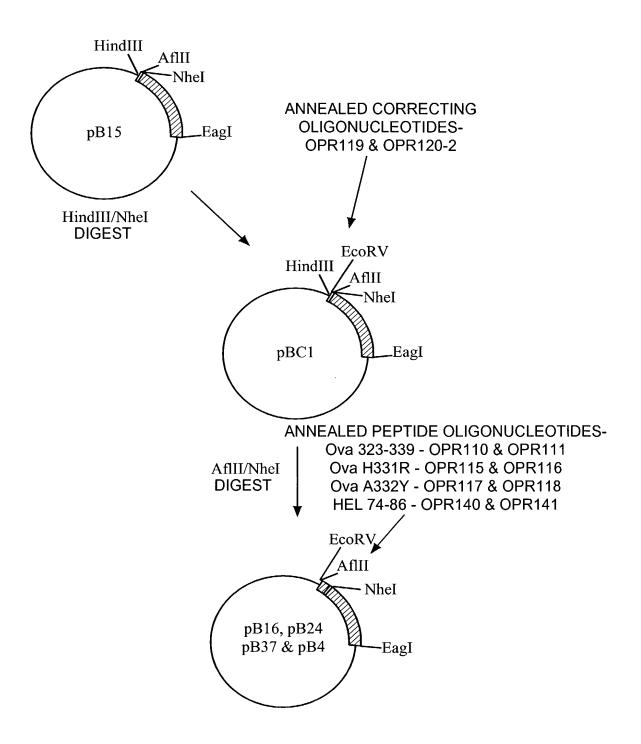


FIG. 3B

### HLA-DRI $\alpha$ CHAIN CLONING SCHEME

6/64

FIG. 4A

TOTAL RNA ISOLATED FROM BLCL K68 CELLS SYNTHESIS OF FIRST STRAND cDNA USING **OLIGO-dT PRIMER** cDNA **TEMPLATE** PCR AMPLIFICATION WITH HindIII DR1A-B & DR1A-F PRIMERS BamHI BamHI HindIII pUC18  $\alpha$ 1 -  $\alpha$ 2-HINGE HindIII/BamHI PCR PRODUCT **DIGEST** HindIII HindIII/BamHI DIGEST K68A3 BamHI TEMPLATE FOR PCR **AMPLIFICATIONS** NcoI PCR WITH AF-N & **AB-S PRIMERS** SpeI FIG. 4A NcoI SpeI pIS139  $\alpha$ 1 -  $\alpha$ 2-HINGE PCR PRODUCT FIG. 4B NcoI/SpeI DIGEST NcoI FIG. 4 -SpeI 39A2

TEMPLATE FOR PCR AMPLIFICATIONS

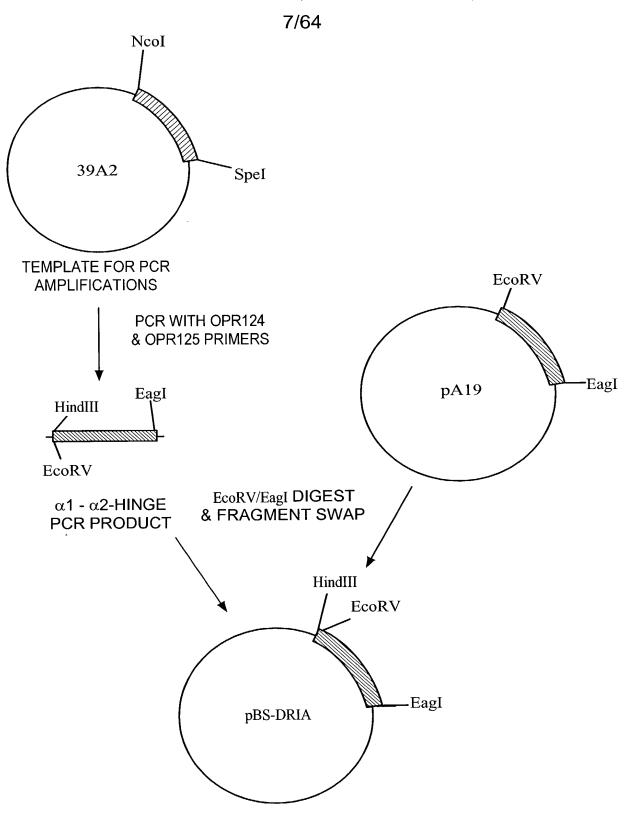
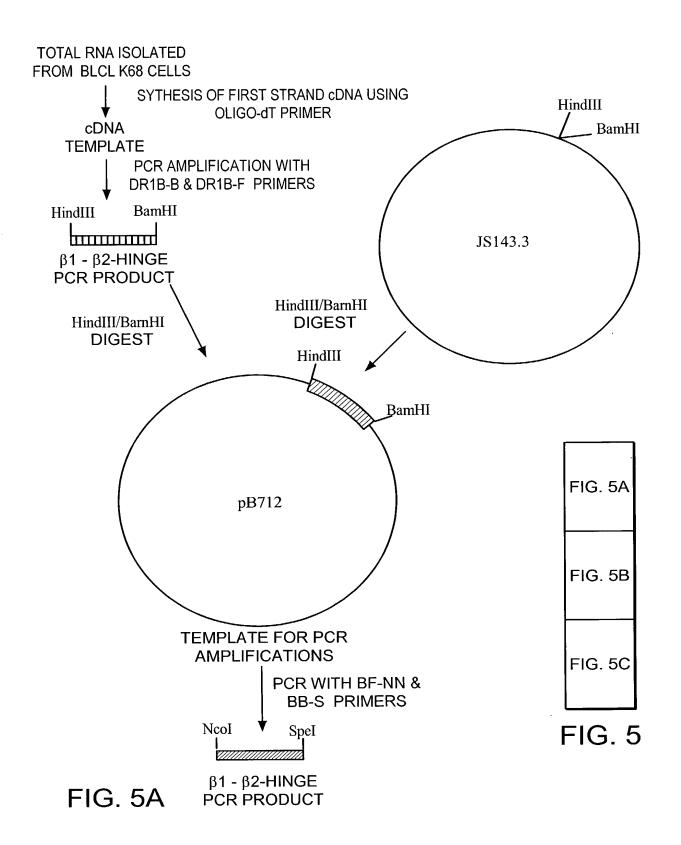


FIG. 4B

8/64 HLA-DRI  $\beta$  CHAIN CLONING SCHEME



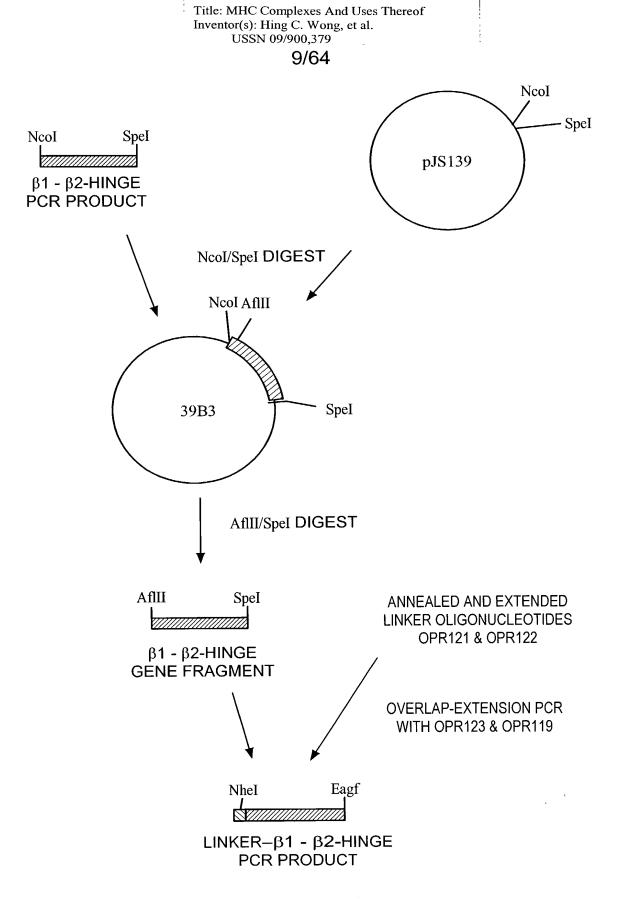


FIG. 5B

FIG. 5C

Nhel

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## I-A $^{\mathbf{S}}$ $\alpha$ CHAIN CLONING SCHEME

TOTAL RNA ISOLATED FROM SJL MOUSE SPLEEN CELLS

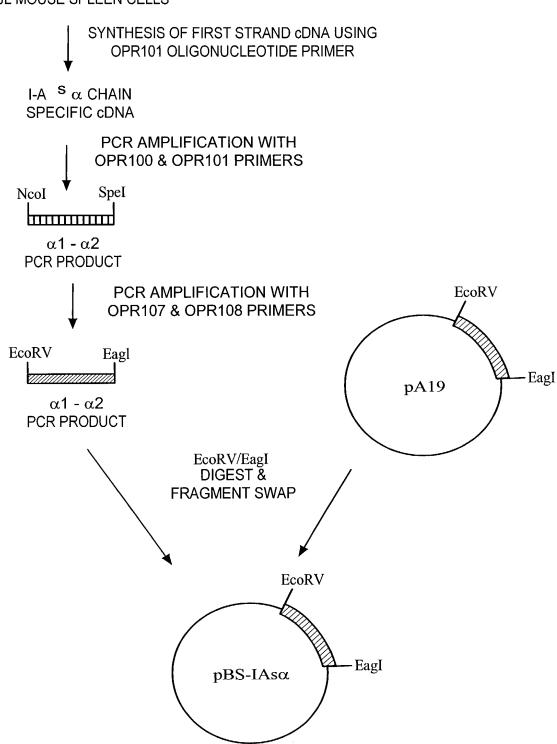


FIG. 6

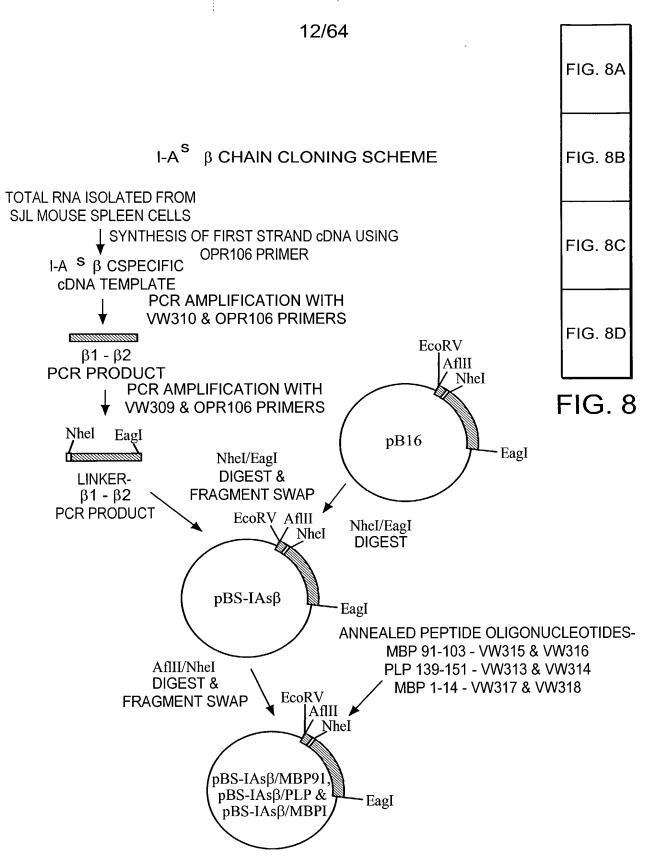


FIG. 7

13/64

# Title: MHC Complexes And Uses Thereof Inventor(s): Hing C. Wong, et al. USSN 09/900,379

#### OLIGOMICLEOTIDES USED IN CONSTRUCTING MHC VECTORS

## I-A<sup>d</sup>/I-A<sup>s</sup> PCR PRIMERS AND CLONING OLIGENUCLEOTIDES (RESTRICTION SITES ARE UNDERLINED)

OPR100

5'-GGG GGG GCC ATG GCC GAA GAC GAC ATT GAG GCC GAC-3'

OPR101

5'-GGG GGG ACT AGT CCA GTG TTT CAG AAC CGG CTC-3'

OPR107

5'-CCC CCC GAT ATC TCA GCT TCC AGC AGT GGA GAC GAC ATT GAG GCC G-3'

OPR108

5'-CCC CCC  $\underline{\text{CGG}}$  CCG CTA CTT ACG TTT CCA GTG TTT CAG AAC CGG C-3'

OPR102

5'-GGG GGG GCC ATG GCC GGA AAC TCC GAA AGG CAT TTC G-3'

OPR104

5'-GCG GCG ACT AGT CCA CTC CAC AGT GAT GGG GC-3'

OPR106

5'-CCC CCC CGG CCG TAC CTG AGG ACC ACT CCA CAG TGA TGG-3'

OPR112

5'-CCC CCC GAT ATC ACA GGT GTC TTA AGT GCT AGC GGA GGG GGC GGA AGC GGC GGA AAC TCC GAA AGG CAT TTC-3'

**OPR119** 

5'-AGC TTG ATA TCA CAG GTG TCT TAA GTG GAG-3'

OPR120-2

5'-CTA GCT CCA CTT AAG ACA CCT GTG ATA TCA-3'

VW310

5'-TCC GGA GGC GGC GGA GAC TCC GAA AGG CAT TTC G-3'

VW309

5'-CGA TCG CTA GCG GCG GTG GTG GTT CCG GTG GCG GCG GAG-3'

**OPR136** 

5'-CCC CCC AGG CTT CCC GGG CCA CCA TGC CGT GCA GCA GAG CTC TG-3'

**OPR139** 

5'-CCC CCC GAG CTC GAA TTC TCA TAA AGG CCC TGG GTG TCT G-3'

OPR132 14/64

5'-CCC CCC AAG CTT CCC GGG CGA CCA TGG CTC TGC AGA TCC CCA GC-3'

**OPR133** 

5'-CCC CCC ACT TAA GGT CCT TGG GCT GCT CAG CAC C-3'

**OPR134** 

5'-CCC CCC CCA TCA CTG TGG AGT GGA GGG-3'

**OPR135** 

5'-CCC CCC GAG CTC GAA TTC TCA CTG CAG GAG CCC TGC TGG-3'

#### HI A-DR1 PCR PRIMERS AND CLONING OLIGONUCLEOTIDES

DR1A-F

5'-GGG GGG AAG CTT ATG ATC AAA GAA GAA CAT GTG ATC ATC-3'

DR1A-B

5'-GCG GCG GGA TCC GTT CTC TGT AGT CTC TGG GAG AGG-3'

DR1B-F

 $5'-GGG \ GGG \ \underline{AAG \ CTT} \ ATG \ GGG \ GAC \ ACC \ CGA \ CCA \ CGT \ TTC \ TTG \ TGG \ CAG \ C-3'$ 

AF-N

5'-GGG GGG GCC ATG GCC ATC AAA GAA GAA CAT GTG ATC ATC-3'

AB-S

5'-GCG GCG ACT AGT GTT CTC TGT AGT CTC TGG GAG AGG-3'

**OPR124** 

5'-GGG GGG <u>AAG CTT GAT ATC</u> TCA GCT TCC AGC AGT AGT ATC AAA GAA GAA CAT GTG ATC-3'

**OPR125** 

5'-GGG GGG CCG CTA CTT ACG TTT CTC TGG GAG AGG GCT TGG AGC-3'

DR1B-B

5'-GCG GCG GGA TCC CTT GCT CTG TGC AGA TTC AGA CC-3'

BF-NN

5'-GGG GGG GCC ATG GCC GGA TCC GCT AGC GGG GAC ACC CGA CCA CGT TTC TTG-3'

BB-5

5'-GCG GCG ACT AGT CTT GCT CTG TGC AGA TTC AGA CCG-3'

#### OPR121

5'-GTT GTC TTA AGT GGA GCT AGC GGA GGG GGC GGG TCC GGA GGT GGT GGG GAC ACC CG-3'

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#### **OPR122**

5'-GAA ATG ACA TTC AAA CTT CAG CTG CCA CAA GAA ACG TGG TCG GGT GTC CCC ACC ACC-3'

#### **OPR123**

5'-GGG GGG CCG TAC CTG AGG ACT TGC TCT GTG CAG ATT CAG-3'

#### PEPTIDE OLIGONUCLEOTIDES.

#### Ova 323-339

### **OPR110**

5'-TTA AGT ATC TCT CAG GCT GTT CAC GCT GCT CAC GCT GAA ATC AAC GAA GCT GGT CGT G-3'

#### OPR111

5'-CTA GCA CGA CCA GCT TCG TTG ATT TCA GCC TGA GCA GCG TGA ACA GCC TGA GAG ATA C-3'

#### Ova H331R

#### **OPR115**

5'- $\underline{\text{TTA}}$  AGT ATC TCT CAG GCT GTT CAC GCT GCT CGG GCT GAA ATC AAC GAA GCT GGT CGT  $\underline{\text{G}}$ -3'

#### OPR116

5'-CTA GCA CGA CCA GCT TCG TTG ATT TCA GCC CGA GCA GCG TGA ACA GCC TGA GAG ATA C-3'

#### Ova A332Y

#### **OPR117**

5'-TTA AGT ATC TCT CAG GCT GTT CAC GCT GCT CAC TAC GAA ATC AAC GAA GCT GGT CGT G-3'

#### **OPR116**

5'- $\underline{\text{CTA}}$  GCA CGA CCA GCT TCG TTG ATT TCA TAG TGA GCA GCG TGA ACA GCC TGA GAG ATA  $\underline{\text{C}}$ -3'

#### HEL 74-86

#### **OPR140**

 $5'-\underline{TTA}$   $\underline{AG}T$   $\underline{AAC}$   $\underline{CTG}$   $\underline{TGC}$   $\underline{AAC}$   $\underline{ATC}$   $\underline{CCC}$   $\underline{TGC}$   $\underline{AGC}$   $\underline{GCC}$   $\underline{CTG}$   $\underline{CTG}$   $\underline{AGC}$ 

#### **OPR141**

5'-CTA GCG GAG CTC AGC AGG GCG CTG CAGBGGG ATG TTG CAC AGG

FIG. 8C

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NP 404-415

OPR129

5'-TTA AGT CAG ATC AGC GTG CAG CCC GCC TTC AGC GTG CAG G-3'

**OPR129** 

5'-CTA GCC TGC ACG CTG AAG GCG GGC TEA ACG CTG ATC TGA C-3'

HA 307-319

**OPR130** 

5'-T  $\overline{TA}$   $\overline{AGT}$   $\overline{CCC}$   $\overline{AAG}$   $\overline{TAC}$   $\overline{GTG}$   $\overline{AAG}$   $\overline{CAG}$   $\overline{AAC}$   $\overline{ACC}$   $\overline{CTG}$   $\overline{AAG}$   $\overline{CTG}$   $\overline{GCC}$   $\overline{ACC}$   $\overline{G-3'}$ 

**OPR131** 

5'-CTA GCG GTG GCC AGC TTC AGG GTG TTC TGC TTC ACG TAC TTG GGA  $\underline{\text{C-3}}$ '

#### MBP 91-103

VW315

 $5'-\overline{\text{TTA}}$  AGT CAC TAT GGC TCC CTG CCG CAG AAG TCC CAG CAC GGG CGC G-3'

VW316

5'-CTA GCG CGC CCG TGC TGG GAC TTC TGC GGC AGG GAG CCA TAG TGA C-3'

#### PLP 139-151

VW313

5'- $\underline{\text{TTA}}$  CAC CAC TCC CTG GGC AAG TGG CTG GGC CAC CCG GAC AAG TTC  $\underline{\text{G-3}}$ '

VW314

 $5\,{}^{\prime}\,{}^{-}\underline{\text{CTA}}$  GCG AAC TTG TTC GGG TGG CCC AGC CAC TTG CCC AGG GAG TGA C-3  ${}^{\prime}$ 

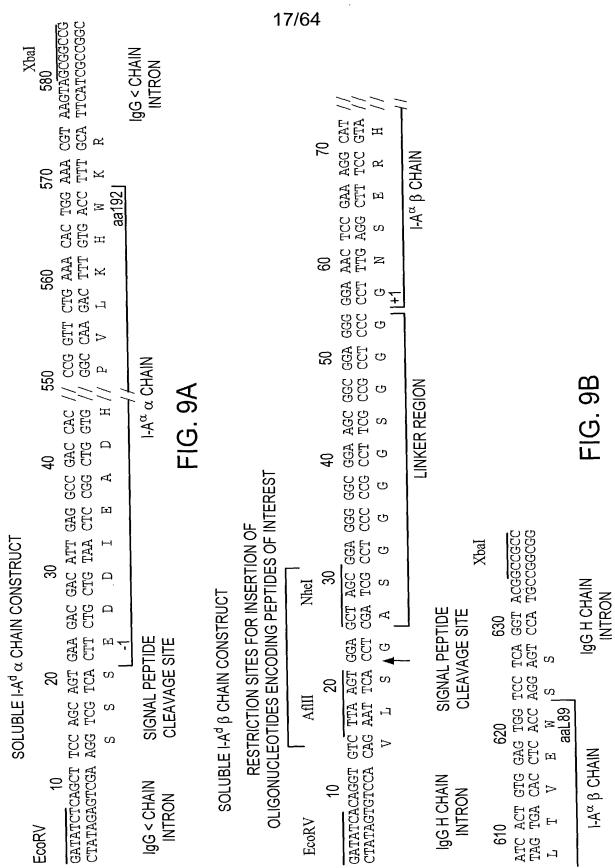
#### MBP 1-14

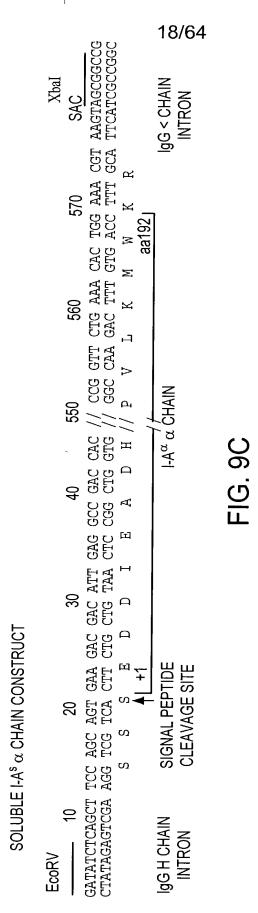
VW317

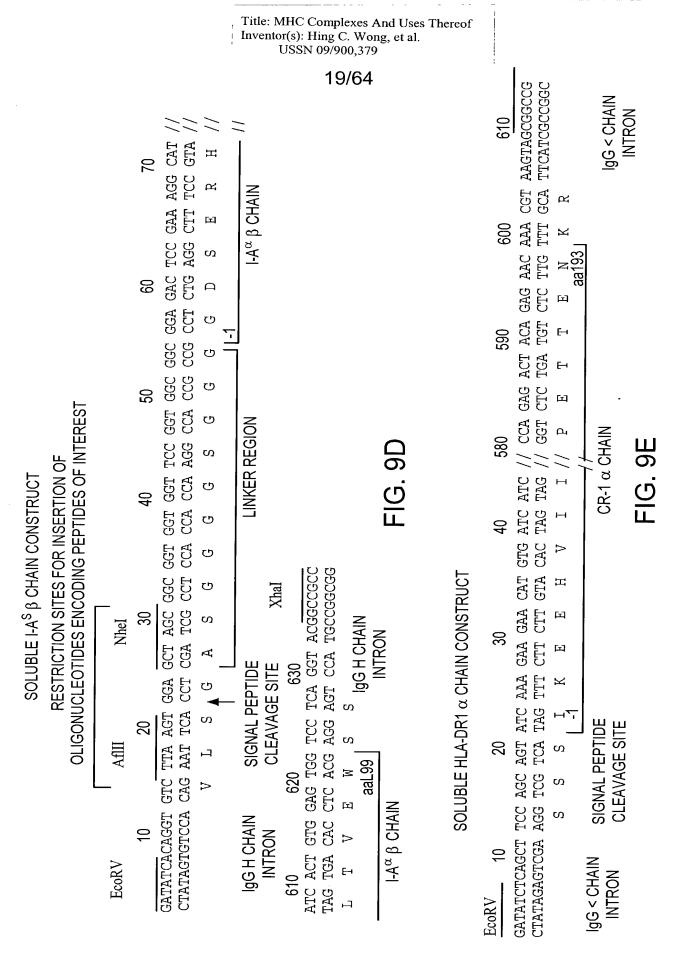
 $5\,\,{}^{\prime}\,-\,T\underline{T}\underline{A}\,\,\underline{A}\underline{G}T$  ATG GCA TCC CAG AAG CGC CCG TCC CAG CGC TCC AAG TAC CTG G-3  ${}^{\prime}$ 

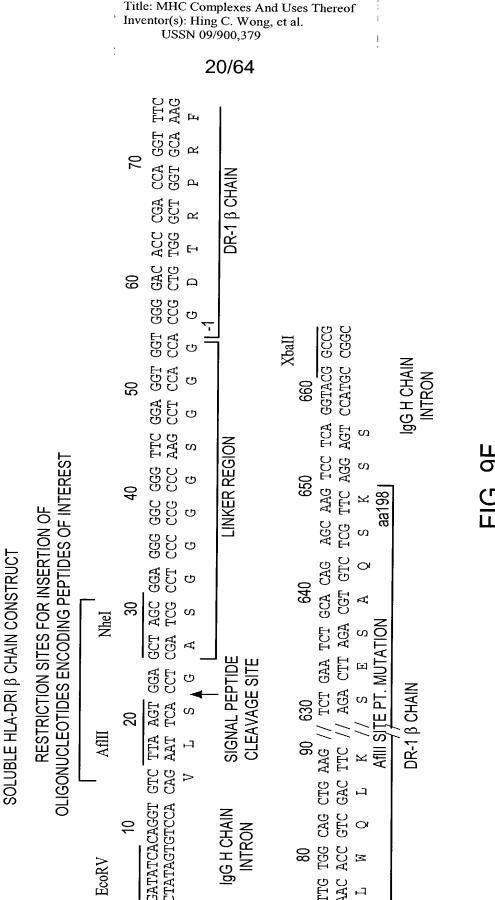
VW316

5'- $\underline{\text{CTA}}$  GCC AGG TAC TTG GAG CGC TGG GAC GGG CGC TTC TGG GAT GCC ATA  $\underline{\text{C}}$ -3'

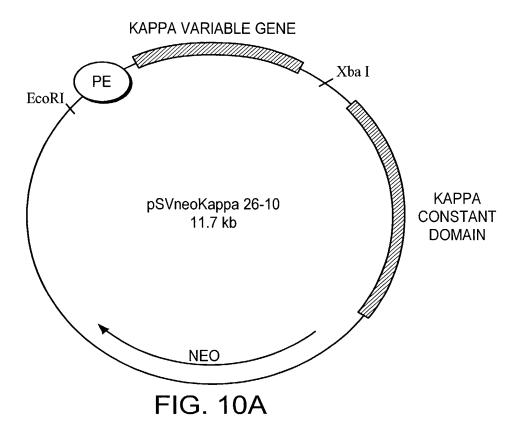


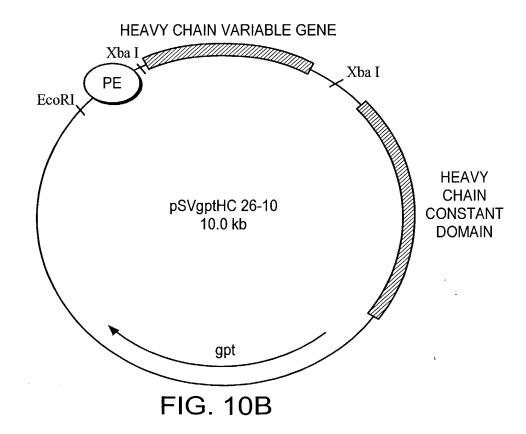




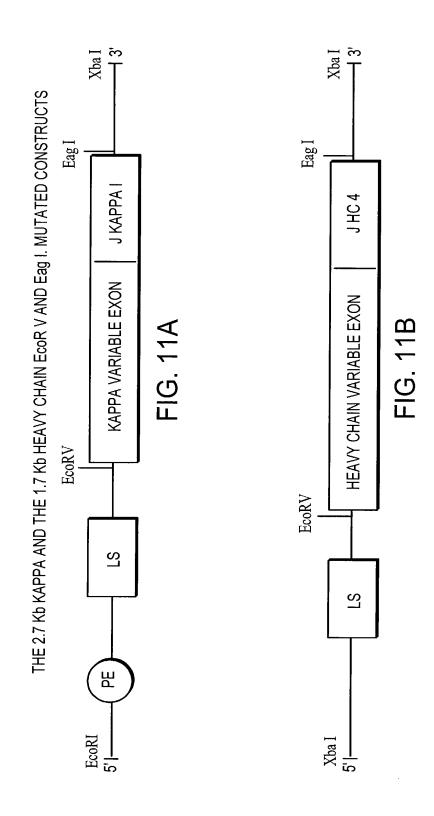


### ORIGINAL MAMMALIAN CELL EXPRESSION VECTORS





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## PCR SITE DIRECTED MUTAGENESIS FOR INTRODUCING EcoRV and Eagl RESTRICTION SITES INTO KAPPA CHAIN 2.7 kb INSERT

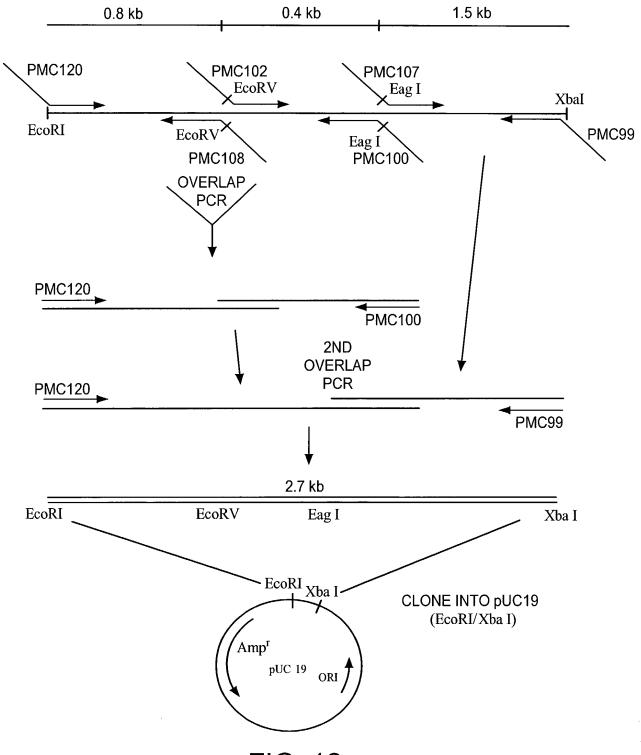


FIG. 12

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## STRATEGY FOR CONSTRUCTING MHC CLASS IIc/KAPPA CONSTANT GENE IN MAMMALIAN CELL EXPRESSION VECTOR

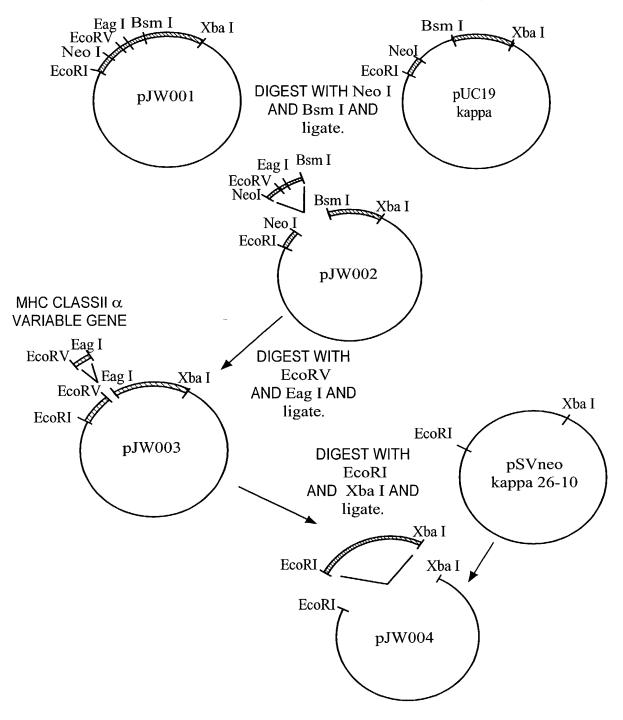


FIG. 13

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PRIMERS USED FOR SEQUENCING MUTATED 2.7Kb FRAGMENT

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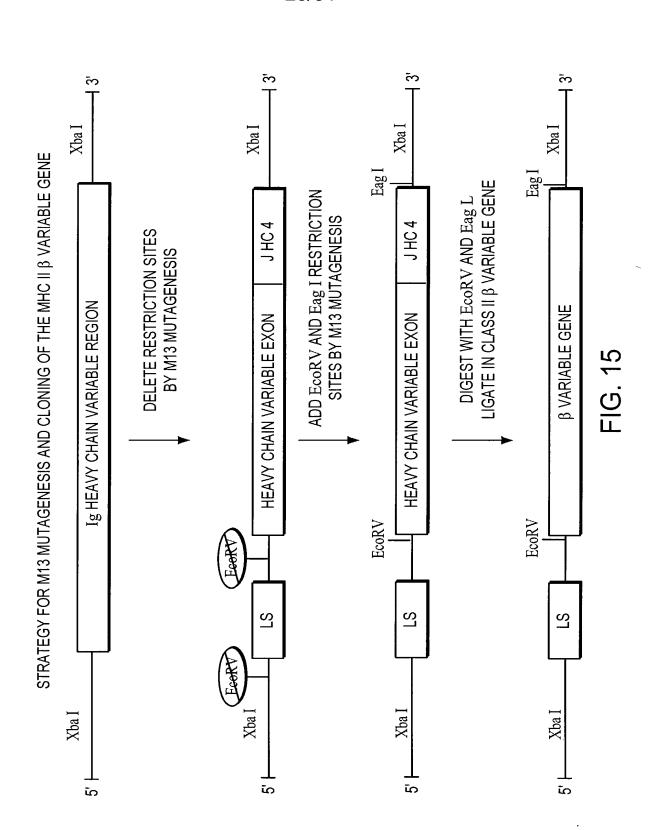
SEQUENCE

PMC-77

(5'GCTTTGCTTACGGAGTTACTC3')

PMC-114

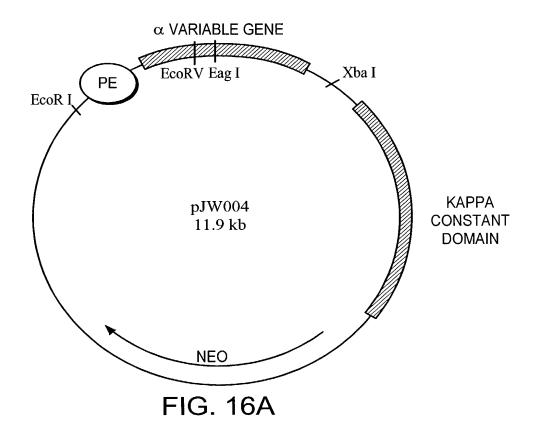
PMC-111

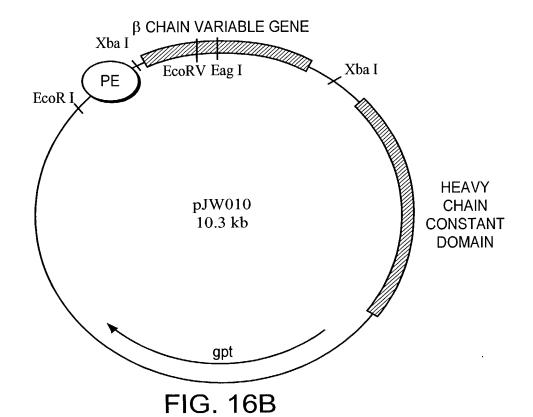


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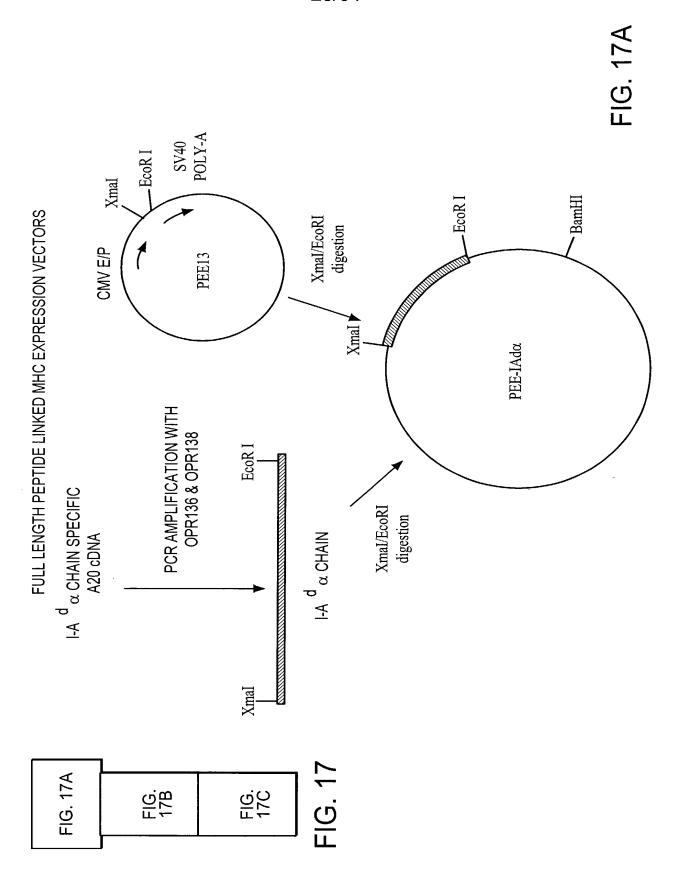
Title: MHC Complexes And Uses Thereof Inventor(s): Hing C. Wong, et al. USSN 09/900,379

FINAL VECTORS FOR EXPRESSING MHC II/Ig CHIMERIC PROTEINS





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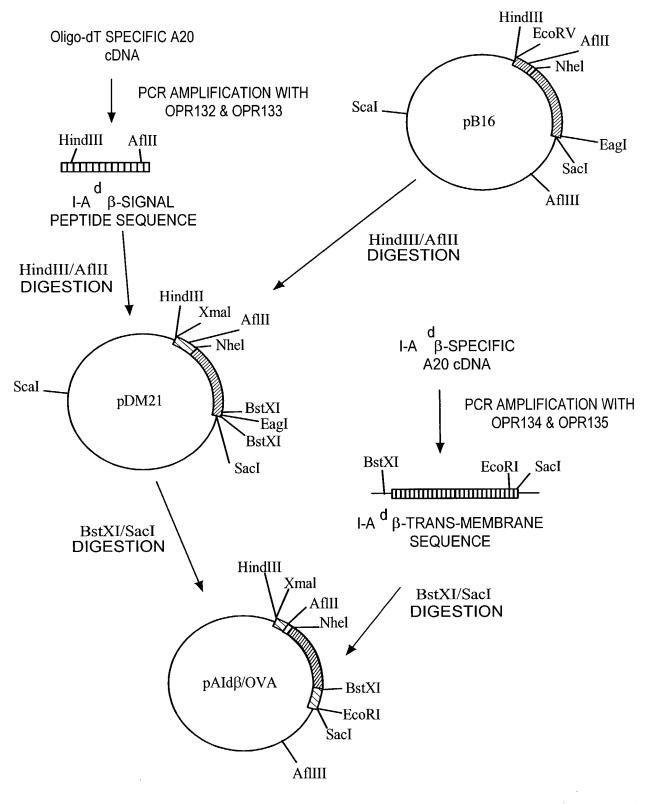


FIG. 17B

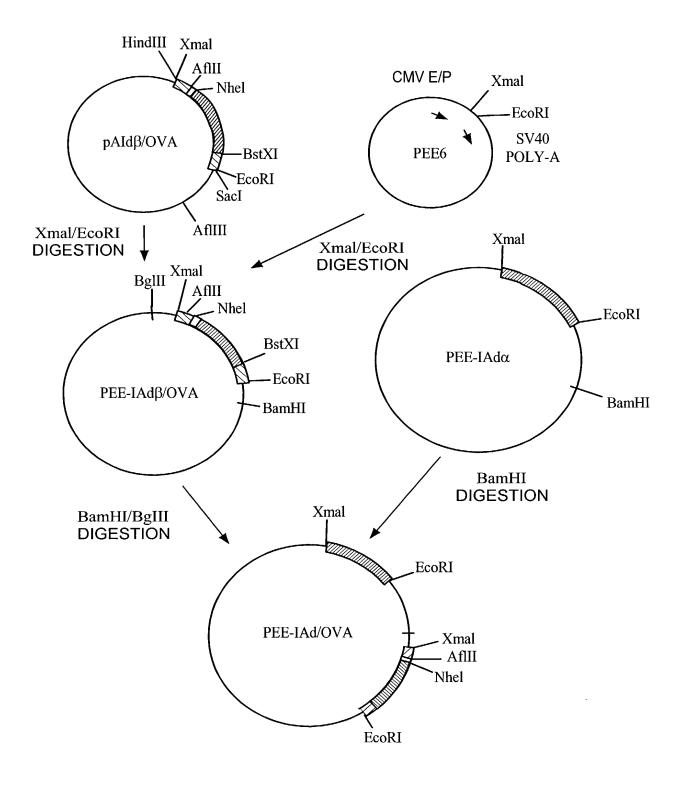
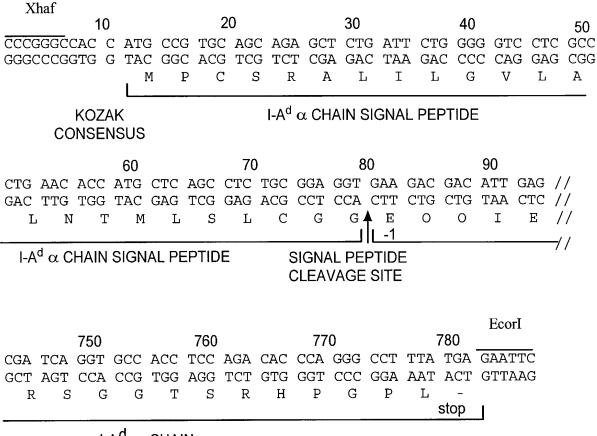


FIG. 17C

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## FULL LENGTH I-A d α CHAIN INSERT



I-A<sup>d</sup>  $\alpha$  CHAIN

FIG. 18A

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## FULL LENGTH I-A<sup>d</sup> β CHAIN INSERT

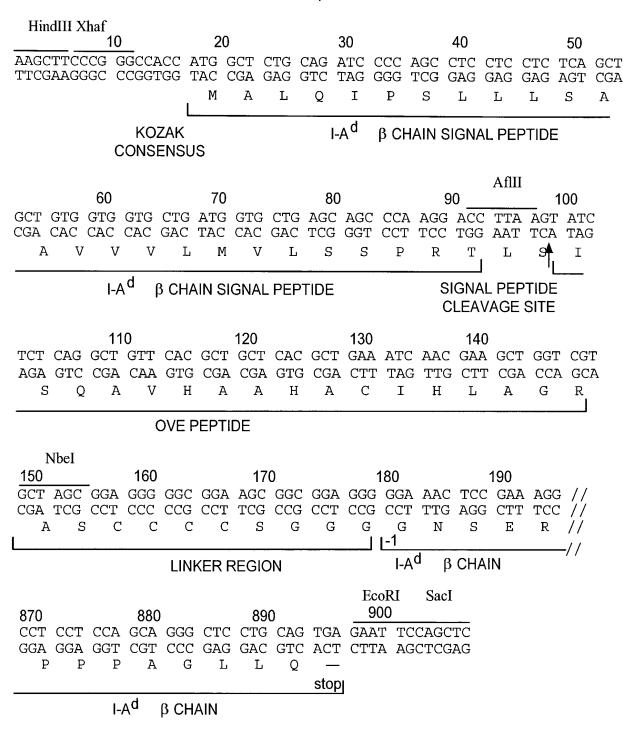


FIG. 18B

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FIG. 19A FIG. 19B FIG. 19C FIG. 19D FIG. 19E FIG. 19F FIG. 19G

FIG. 19

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### FULL-LENGTH PEPTIDE LINKED MHC EXPRESSION VECTORS

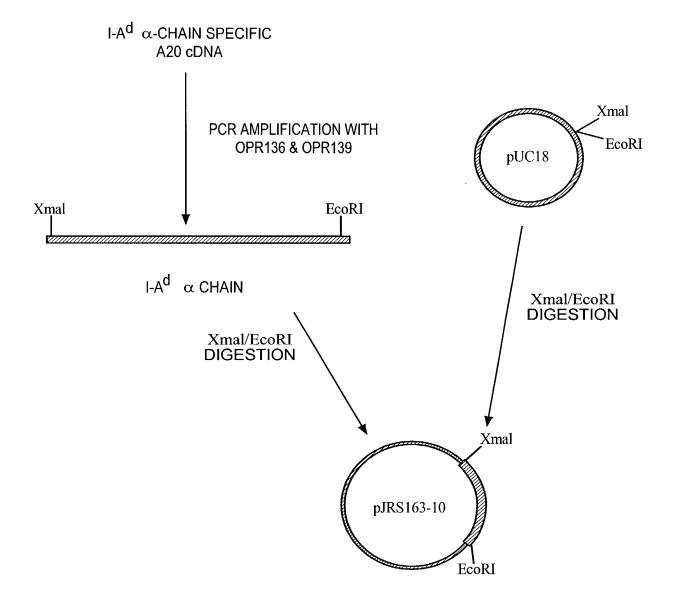


FIG. 19A

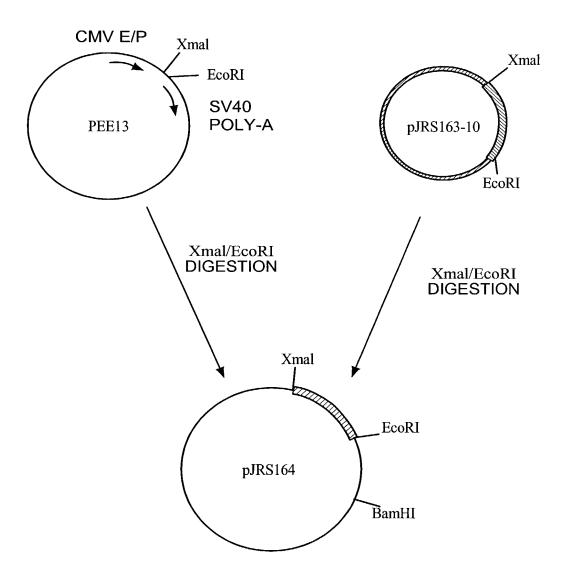


FIG. 19B

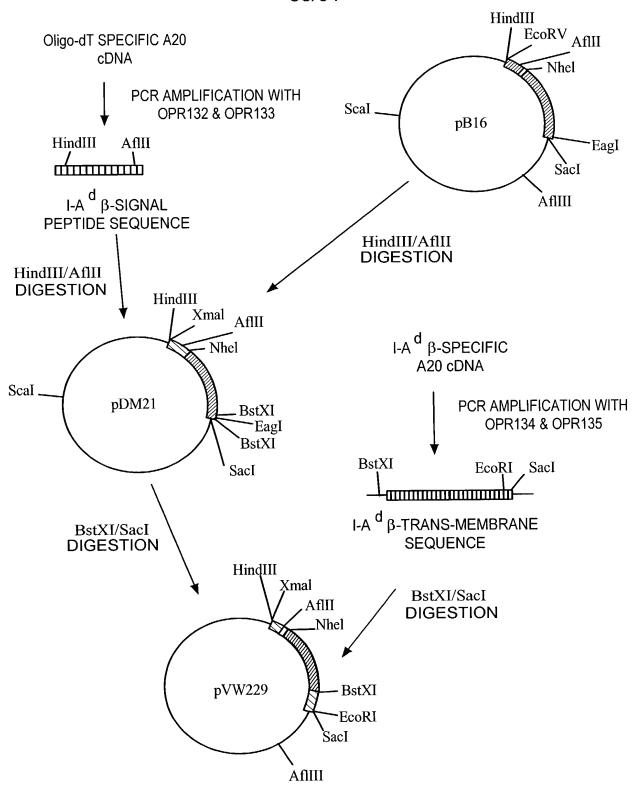


FIG. 19C

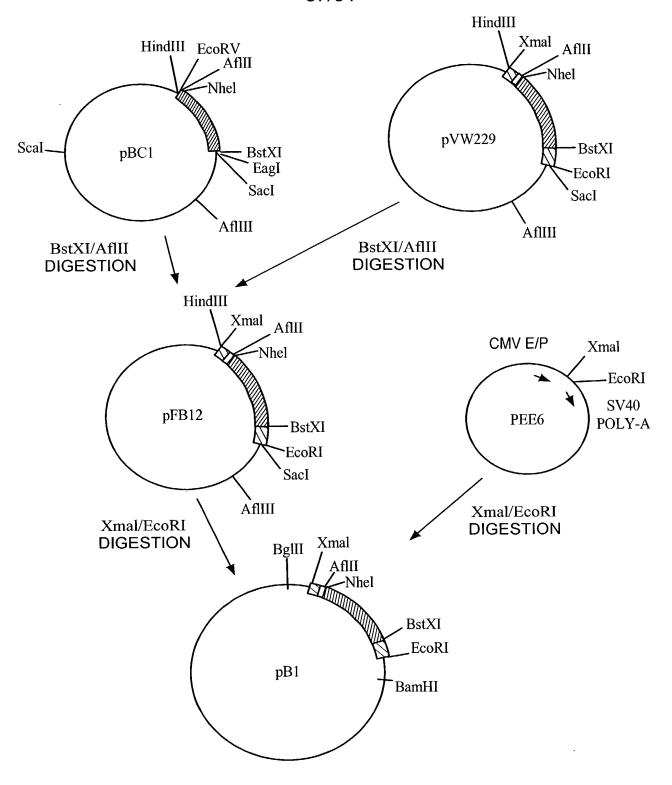


FIG. 19D

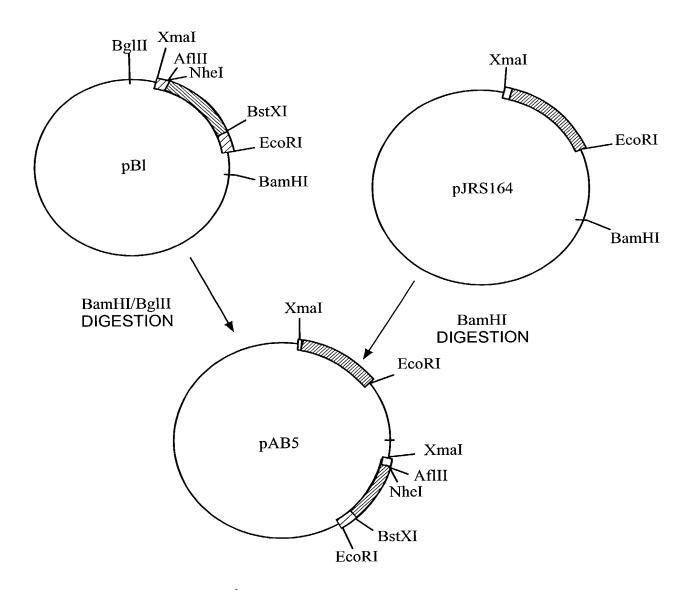


FIG. 19E

Inventor(s): Hing C. Wong, et al. USSN 09/900,379 39/64 HindIII Xmal AflII HindIII EcoRV AflII Nhel -Nhel pVW229 ScaI. -BstXI pB4 EagI EcoRI SacI SacI AflIII ÀfIII BstXI/AflII BstXI/AflII **DIGESTION DIGESTION** HindIII Xmal AfIII CMV E/P Xmal -Nhel -EcoRI SV40 pFBH3 PEE6 POLY-A BstXI EcoRI SacI AfIIII Xmal/EcoRI Xmal/EcoRI **DIGESTION DIGESTION** Xmal BglII AflII -Nhel BstXI \_EcoRI pBH4 BamHI

Title: MHC Complexes And Uses Thereof

FIG. 19F

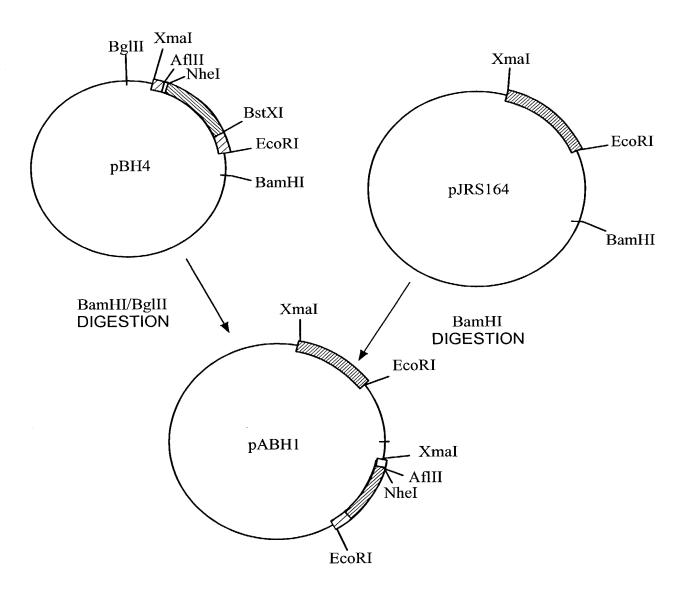


FIG. 19G

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#### OLIGONUCLEOTIDES USED IN CLONING

#### **OPR132**

I-A  $^d$   $_\beta$  SIGNAL PEPTIDE FRONT PRIMER WITH KOZAK CONSENSUS FOR CellTech VECTOR-HindIII/Xmal SITES 5' -CCC CCC AAG CTT CCC GGG CCA CCA TGG CTC TGC AGA TCC CCA GC-3'

#### **OPR133**

I-A  $^d$   $\beta$  SIGNAL PEPTIDE BACK PRIMER WITH KOZAK CONSENSUS FOR CellTech VECTOR- AfIII SITE 5' -CCC CCC ACT TAA GGT CCT TGG GCT CAG CAC C-3'

#### **OPR134**

I-A  $^d$   $\beta$  TRANSMEMBRANE FRONT PRIMER FOR CellTech VECTOR- BstXI SITES 5' -CCC CCC CCA TCA CTG TGG AGT GGA GGG-3'

#### **OPR135**

I-A  $^d$   $\beta$  TRANSMEMBRANE BACK PRIMER FOR CellTech VECTOR- SstI, EcoRI SITES 5' -CCC CCC GAG CTC GAA TCC TCA CTG CAG GAG CCC TGC TGG-3'

#### **OPR136**

I-A<sup>d</sup> a SIGNAL PEPTIDE FRONT PRIMER WITH KOZAK CONSENSUS FOR CellTech VECTOR-HindIII/Xmal SITES
5' -CCC CCC AAG CTT CCC GGG CCA CCA TGC CGT GCA GAG CTC TG-3'

#### **OPR139**

I-A  $^d$   $\alpha$  TRANSMEMBRANE PRIMER FOR CellTech VECTOR- SstI/EcoRI SITES 5' -CCC CCC GAG CTC GAA TCC TCA TAA AGG CCC TGG GTG TCT G-3'

#### B7-1-2F

MURINE B7-1 FRONT PRIMER WITH KOZAK CONSENSUS FOR CellTech VECTOR- Not1 SITES 5' - CCC CCC CCG CGG CCG CCC CAC CAT GGG ACT GAG TAA CAT TCT C-3'

#### B7-1-2B

MURINE B7-1 BACK PRIMER FOR CellTech VECTOR- Notl SITE 5' - CCC CCC GCG GCC GCT TTA AAA ACA TGT ATC ACT TTT-3'

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# NSO/ClassII/OVA CLONES STIMULATE IL-2 PRODUCTION FROM DO11.10

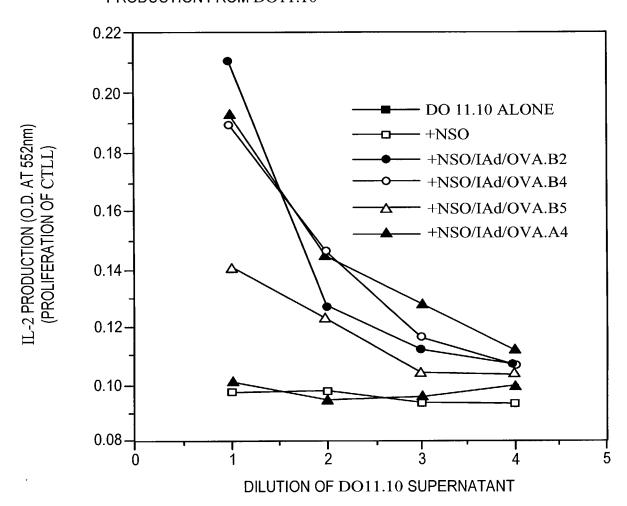


FIG. 21



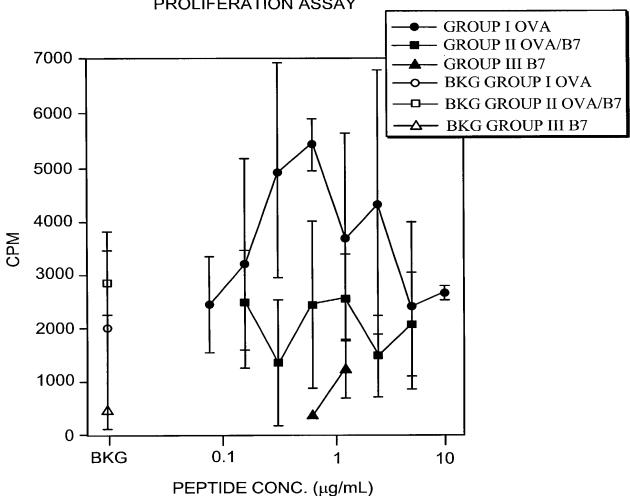


FIG. 22

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# INTRADERMAL IAd/OVA & IAd/HEL DNA INJECTIONS PROLIFERATION ASSAY

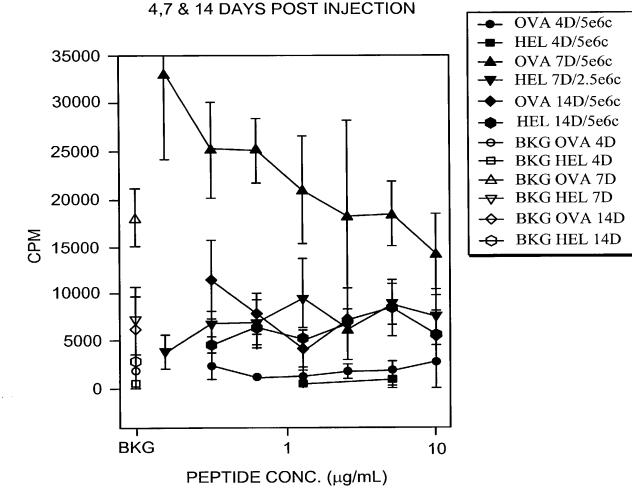
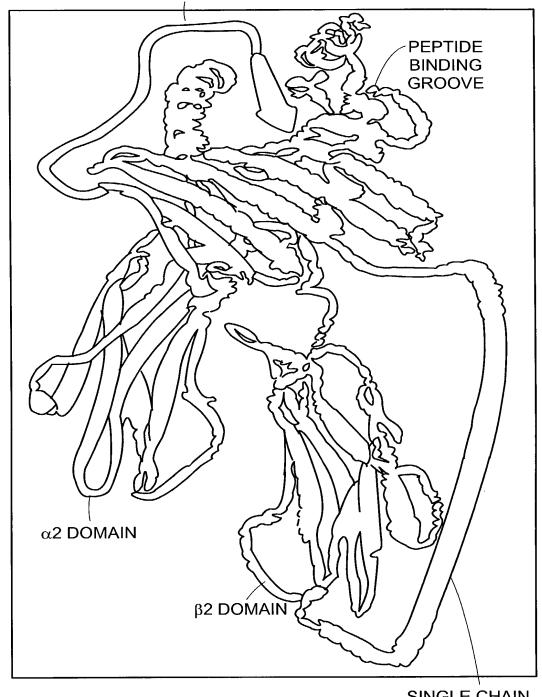


FIG. 23

45/64 LINKER SEQUENCE LINKED TO PRESENTING PEPTIDE



SINGLE CHAIN LINKER SEQUENCE

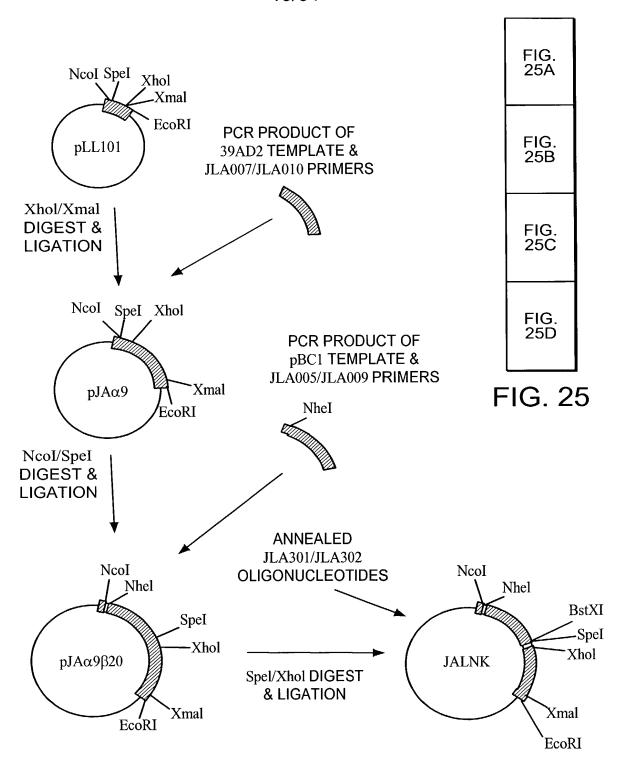
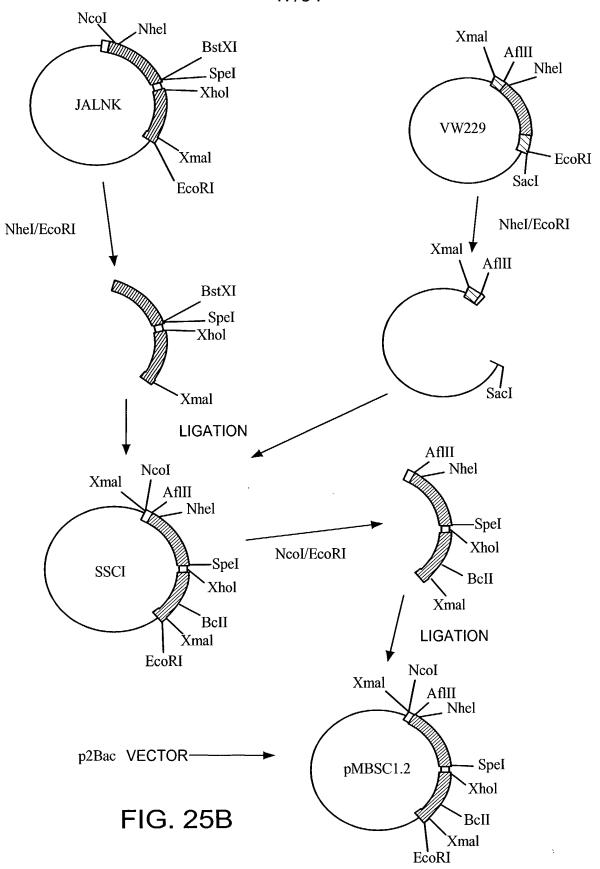


FIG. 25A



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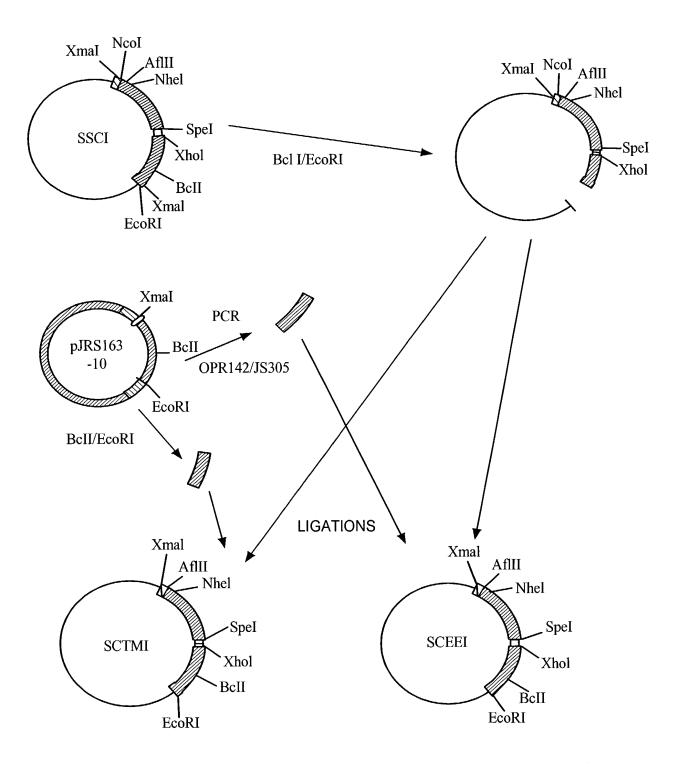


FIG. 25C

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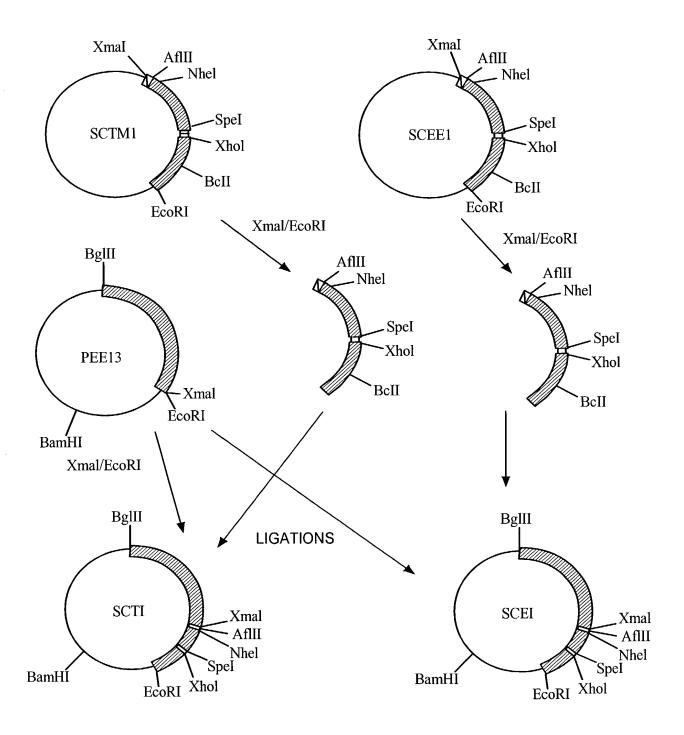


FIG. 25D

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#### JLA-005

5' - CCCCCGCCATGGCCGCTAGCGGAGGGGGGGGGAAGC-3'

#### JLA-007

5' -CCCGGGGCCTCGAGTGAAGACGACATTGAGGCCGAC-3'

#### JLA-009

5' -CCCCCCACTAGTCCACTCCACAGTGATGGGGCT-3'

#### JLA-010

5' - CCCCCCCCGGGACCAGTGTTTCAGAACCGGCTCCTC-3'

#### JLA-301

5' -TCGAGGAACCGCCACCGCCAGAACCGCCGCCACCGGA-ACCACCACCGCCGCTGCCACCGCCACCA-3'

#### JLA-302

5 ' - CTAGTGGTGGCGGTGGCAGCGGCGGTGGTTCCGG-TGGCGGCGGTTCTGGCGGTGGCGGTTCC-3 '

#### **OPR-142**

5' - CTTGGGAATCTTGACTAAGAGG-3'

#### JS-305

5' - CAGGTCGAATTCTCATTCCATCGGCATGTACTCTTCTT - CCTCCCAGTGTTTCAGAACCGG - 3'

FIG. 26

FIG. 27A	
FIG. 27B	
FIG. 27C	
FIG. 27D	

FIG. 27

# FIG. 27A

-1G. Z/A		51/64			
10		20	30	40	50 *
CCACC ATG GCT	A GAC GTC TA		AG GAG GAG	AGT CGA CG	A CAC CAC
6	0	70 *	80 *	90 *	
GTG CTG ATG ( CAC GAC TAC ( V L M	CAC GAC TCG V L S	AGC CCA AGG	TGG AAT T	CA TAG AGA S I S	
100		120			
GTT CAC GCT (CAA GTG CGA (CAA GTG CGA (CAA GTG CGA (CAA GTG CGA (CAA GTG CAA G	GCT CAC GCT	GAA ATC AAC	GAA GCT G CTT CGA C	GT CGT GCT CA GCA CGA	TCG CCT
	160	170 *	180		
GGG GGC GGA A CCC CCG CCT T G G G 10 AMINO	AGC GGC GGA FCG CCG CCT S G G	CCC CCT TTG	TCC GAA A AGG CTT T S E	GG CAT TTC CC GTA AAG R H F	GTG GTC CAC CAG V V>
200	210	220	2	230	240
CAG TTC AAG ( GTC AAG TTC ( Q F K	GGC GAG TGC CCG CTC ACG	TAC TAC ACC ATG ATG TGG	AAC GGG A TTG CCC T	CG CAG CGC GC GTC GCG	ATA CGG TAT GCC
250	260 *		70 *	280	290
CTC GTG ACC A	TCT ATG TAG	ATG TTG GCC Y N R	CTC CTC A	TG CAC GCG	ATG CTG Y D>
30		310	320	330	
AGC GAC GTG (TCG CTG CAC S D V	GGC GAG TAC CCG CTC ATG	CGC GCG GTC GCG CGC CAC	ACC GAG C TGG CTC G	TTG GGG CGG AC CCC GCC	GGT CTG
340	350 *	360	370	380	
			**	••	CGG GCC

						SSN 0	ng C. 9/900,	,379				ì			
Э. 2	27E	3					52/6	4							
39	0		4	00			41(			42			4	30	
GAG CTC	GTG CAC	CTG	TGC	GCG CGC	ACG C	TCT R	CAC GTG H	AAC TTG N	ATG Y	GAG CTC	GGG CCC G	GGC	CTC	ACC	
	440	)		45	0			460 *			470 *			48	0
Τ	AGG S	GAC L	GCC R	CGG GCC R	CTT GAA L	CTT E	CAG GTC O	CCC GGG P	AAT TTA N	GTC CAG V		TAG I	AGG S	CTG GAC L	TCC AGC S>
	4	190 *			50¢			5	10 *		5	20 *			53
	TGT	CTC	CGG	GAG	TTG	GTG	GTG	TTG	TGA	GAC	GTC CAG V	ACA	AGC	CAC	TG
		:	540			550 *			560 *			57 *			
CTA	AAG	ATG Y	GGT P	CGG A	TTC K	ATC TAG I	TTT K	CAC V	CGC GCG R	TGG ACC W	TTC AAG F	AGG TCC R	AAT TTA N		
580			590	)						610 *			620		
GAG		TGT	GTG CAC	GGG	CAG	AGT	TCC AGG	TGT	CAG GTC	CTT GAA	ATT TAA I	TCC	AAT TTA		CT
	30		(	640			650			66	50		6	70 *	
TGG ACC W	TGG T	AAG F	GTC Q	CAG	GAC L	CAG	TAC	CTG GAC	CTC	TAC	ACC TGG T	GGA	GTA	CAG GTC	
	680				90			700 *			71	0		72	20
CTC	GTC CAG	ATG V	TGG T	ACG C	GTA H	CAC V	CTC E	GTA H	GGG P	TCG S	CTG GAC L 3-2 D	TTC A	TCG S	GGG P	TA I>
	7:	30			740	)		75			-				770
TCA	CAC	CTC	ACC	TGA	AGT TCA S	GGT CCA G	CCG G	GGT CCA G	GGC CCG G	TCG S	GGC CCG G ACIE	GGT CCA G	CCA G	CCA G	TC AG S>

# FIG. 27C

	<b>27</b> (							•							
		7	80		•	790 *			80	0		8	10 *		
CCA G	GGC CCG G	CCG G	CCA G	AGA S	CCG G	CCA G	CCG G	CCA G	AGG S	AGC S	TCA S	CTT E	CTG D	CTG D	TAA I>
				)		8			8				860 *	)	
CTC E	GCC CGG A I-A	CTG D	GTG H	CAT V	CCG G	AAG F	ATA Y	CCA G	TGT T	TGA T	CAA V	ATA Y	GTC Q	AGA S	
	1- <i>7</i> 870 *	Au α-		880 *				90							
CCT G	GAC CTG D	TAA I	CCG G	GTC Q	ATG Y	TGT T	GTA H	CTT E	AAA F	CTA D	CCA G	CTA D	CTC E	AAC L	AAG
	920			9				940	)			0		9	60
ATA	GTG CAC V	CTG	AAC	CTA	TTC	TTC	TTT	TGA	CAG	ACC	TCC	GAA	GGA	CTC	AAA
	9	70 *			980			9	90		1	000			1010
CCG	CAA GTT Q	AAC	TAT	GAG	AAA	GAG CTC	CCC GGG	CAA GTT	GGT CCA	GGA CCT	CTG GAC	CAA GTT	AAC TTG	ATA TAT	GCT
		10	20			1030			10	40		10	050		
CGT	CTT	TTT	CAC GTG	TTG	AAC	GGA CCT	ATC TAG	AAC	TGA	TTC	TCC	AGT	TTA	AAG	ACC TGG T>
									1-/	au α	-100	>1417 (11			
1060						1	080				)			0	
1060 * CCA GGT P		ACC TGG T	1070 * AAT TTA N	GAG CTC E	GCT CGA A	1 CCT GGA P	080 * CAA GTT Q	GCG CGC A	ACT TGA T	1090 * GTG CAC V	TTC AAG F	CCC GGG P	110 * AAG TTC K	TCC AGG S	CCT GGA P>
1060 * CCA GGT P	GCT CGA A	ACC TGG T	1070 * AAT TTA N ><	GAG CTC E	GCT CGA A <b>I</b> -A	1 CCT GGA P Ad α	080 * CAA GTT Q -2 DC	GCG CGC A	ACT TGA T	109( GTG CAC V	TTC AAG F	CCC GGG P	AAG TTC K	TCC AGG S	CCT GGA P>

		1160 *	)		117	70		•	1180	)		119	0		12	00
	TTC AAG	CCA GGT	GCT GGA	GTG CAC	ATC TAG	AAC TTG	ATC TAG	ACA TGT	TGG ACG	CTC GAG	AGA TCT	AAT TTA	AGC TCG	AAG TTC K	TCA AGT	GTC CAG
		12	210 *		1	220			12	:30		1	240			1250
	TGT	GAC CTG	GGC CCG	GTT CAA	TAT ATA	GAG CTC	ACC TGG	AGC TCG	TTC AAG	CTC GAG	GTC CAG	AAC TTG	CGT GCA	GAC CTG D	CAT GTA	TCC AGG
														: :90 *		
	AAG	GTG	AAG TTC	CTG GAC	TCT AGA	TAT ATA	CTC GAG	ACC TGG	TTC AAG	ATC TAG	CCT GGA	TCT AGA	GAT CTA	GAT CTA D	CTG	TAA
1:	300 *			1310 *	)		1:	320			1330			1340	)	
	ATA	CTG D	ACG C	TTC K	CAC V	CTC E	GTG H	ACC W	CCG G	GAC L	CTC E	CTC E	GGC P	GTT CAA V	GAC` L	TTT K>
	125												MAIN	<b>\</b>		
	130	*		13	*			137	U		13	*				
	GTG H	ACC W	AGG S	GCC	CGA A	TCA S	GTG H	GTA H	GTG H	GTA H	GTA H	CAC GTG H	ATC			

FIG. 27D

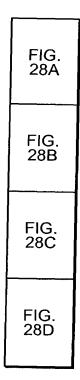


FIG. 28

S D V

G E

Y R

#### 56/64

FI	G.	28	3A					36/6	) <del>4</del>							
			10	)			20			30 *			40 *			50 *
	CCAC GGT(	GG TA	AC CO	GA GA	AC G	rc T	AG G	GG T	CG G	AG G.	AG G.	AG A	GT C	GA C	'GA C	TG GTG CAC CAC V V>
			(	60 *			70 *			80			9			
	CAC	GAC L	TAC M	GTG CAC V	GAC L	TCG S	TCG S	GGT P	TCC R	ACC TGG T	AAT L	TCA S	TAG I	AGA S	GTC Q	CGA A>
	100			110	)		12	0		,	130			140		
	'AA (	TG (	CGA	CGA (	CAC (	GCT CGA	GAA CTT	ATC TAG	TTG	GAA CTT E	CGA	CCA	GCA	CGA	TCG	CCT
	1	50 *			160 *			170			18	30 *			90	
	CCC	CCG	CCT	AGC TCG	GGC	GGA	GGG	GGA CCT	AAC TTC	TCC AGG	CTI	TCC	: GTA	AA(	G CAC	G GTC C CAG V>
		20	0		21	10		2	20			230	)		24	
_	GTC	TTC AAG	AAG TTC	CCG	CTC	ACG	TAC ATG	TAC ATG	ACC TGG	AAC TTG N	CCC	TGC	GTC	GCG	TAT	CGG GCC R>
		2	250			260			27	70 *		2	80			290
	CTC GAG L	GTG CAC V	ACC TGG	AGA TCT R	TAC ATG Y	ATC TAG	TAC	TTG	CGG GCC	GAG CTC	GAG CTC E	TAC	GTG	CGC GCG R	TAC ATG Y	GAC
			30	00		3	310			320			33			
	AGC	GAC	GTG							ACC						GAC

350 360 370 380 340 GCC GAG TAC TGG AAC AGC CAG CCG GAG ATC CTG GAG CGA ACG CGG GCC CGG CTC ATG ACC TTG TCG GTC GGC CTC TAG GAC CTC GCT TGC GCC CGG Y S Q Ρ E Ι m L $\mathbf{E}$ 

TCG CTG CAC CCG CTC ATG GCG CGC CAC TGG CTC GAC CCC GCC GGT CTG

T  $\mathbf{E}$  L G

R P D>

A V

FIC	G.	28	В					57/6	64							
	3	390 *			400				10 *		4	20		•	430 *	
		CAC	CTG	TGC	CGC	ACG	TCT	GTG	TTG	ATG	CTC	GGG CCC G	GGC	CTC	ACC	TCG
		44(	0									470	)		48	0
	TGG	AGG	${\tt GAC}$	GCC	CGG GCC	GAA	CTT	$\operatorname{GTC}$	GGG	TTA	CAG	GCC CGG A OMAI	TAG	AGG	GAC	AGG
			490			500						5				530
		TGT	CTC	CGG	GAG	TTG	GTG	GTG	AAC TTG	TGA	GAC	GTC CAG V	ACA	AGC	CAC	TGT
				540			550			560			57			
	CTA	AAG	ATG	GGT	CGG	TTC	TAG	TTT	CAC	GCG	TGG ACC	TTC AAG F	TCC	TTA	CCG	GTC
5	580			590			60	0		6	10			620 *		
	GAG		TGT	CAC	CCC	CAG	AGT	AGG	TGT	GTC	CTT GAA	ATT TAA I	TCC	AAT TTA		CTG
	6	30		6	640 *			650 *			66	0		6	570 *	
		ACC TGG	AAG		GTC CAG	GAC L	CAG	ATG TAC M	GAC L	CTC E	TAC M	TGG T	GGA		CAG GTC	GGA CCT G>
		 68(			69			7	00			710			720	) *
	CTC	GTC CAG	TAC ATG	TGG	TGC ACG	CAT GTA	CAC	GAG CTC	CAT GTA	GGG	TCG	CTG GAC L MAIN	TTC	TCG	CCC GGG	ATC TAG
			730			740				50		7	60			770
	TGA	CAC	CTC	ACC	TGA	TCA	GGT CCA	CCG	CCA	CCG	TCG	GGC CCG G IINO A	CCA	CCA	CCA	AGG

FIG. 28C

	J.	28						30/1	<b>J</b> 1							
			78	3 <b>0</b>		7	790			800	)		81	0		
C	CA	CCG	CCG	CCA	AGA S	CCG G	CCA G	CCG G	CCA G	TCC AGG S	AGC S	TCA S	CTT E	GAC CTG D	GAC CTG D	TAA I>
82	0			830			84				50 *			860		
C	TC.	CGG	CTG	GTG	CAT	CCG	AAG	ATA	CCA	TGT	TGA	CAA	АТА	GTC	TCT AGA S	GGA
	87				80				)		90			9		
С	CT	GAC CTG	TAA	CCG	CAG GTC	ATG	ACA TGT	CAT GTA	GAA CTT	AAA		GGT CCA	CTA	CTC	TTG AAC	AAG
		920			93	30 *		9	40		·	 950			96	0
	TΑ	CAC	CTG	AAC	CTA	AAG TTC	AAG TTC	AAA TTT	ACT TGA	GTC CAG	TGG ACC	AGG TCC	CTT GAA	GGA	GAG CTC E	AAA
		9	70			980			go	Ω		10	00		4	010
			*						55			10			ı	010
С	CG	GTT	* TTG AAC	TAT	CTC GAG	* TTT AAA	GAG CTC	CCC GGG	CAA GTT	* GGT CCA	GGA CCT	CTG GAC	* CAA GTT	AAC TTG	ATA TAT I	* GCT CGA
С	CG	GTT	* TTG AAC L	TAT I	CTC GAG L	TTT AAA F	GAG CTC E	CCC GGG P	CAA GTT Q	* GGT CCA G	GGA CCT G	CTG GAC L	* CAA GTT Q	AAC TTG N	ATA TAT	* GCT CGA
G 	CG G 	GTT Q GAA CTT	TTG AAC L 102 AAA TTT	TAT I 20 * CAC GTG	CTC GAG L AAC	TTT AAA F  10  TTG AAC	GAG CTC E  O30 * GGA CCT	CCC GGG P ATC TAG	CAA GTT Q 1 TTG	GGT CCA G VALUE ACT TGA	GGA CCT G AAG	CTG GAC L AGG	CAA GTT Q 105	AAC TTG N 50 * AAT	ATA TAT I TTC AAG	GCT CGA A>
G G C	CG G CA GT A	GTT Q GAA CTT	TTG AAC L 102 AAA TTT K	TAT I 20 * CAC GTG H	CTC GAG L AAC TTG N	TTT AAA F  10  TTG AAC L	GAG CTC E 	CCC GGG P ATC TAG I	CAA GTT Q 1 TTG AAC L	GGT CCA G O40 * ACT TGA	GGA CCT G AAG TTC K	CTG GAC L AGG TCC R	CAA GTT Q 105 TCA AGT S	AAC TTG N 50 * AAT TTA N OMAI	ATA TAT I TTC	GCT CGA A>
G G G G G G G G G G G G G G G G G G G	CG G  CA CGT A 	GAA CTT E	TTG AAC L 102 AAA TTT K ACC TGG	TAT I 20 * CAC GTG H 1070 * AAT TTA	CTC GAG L AAC TTG N	TTT AAA F  10  TTG AAC L  GCT CGA	GAG CTC E D30 * GGA CCT G	CCC GGG P ATC TAG I * CAA GTT	CAA GTT Q 1 TTG AAC L	GGT CCA G 040 * ACT TGA T 10 ACT TGA	GGA CCT G AAG TTC K 	CTG GAC L AGG TCC R -Ad c	* CAA GTT Q 105 TCA AGT S c-1 DC	AAC TTG N  50 * AAT TTA N OMAI  4 AAG TTC	ATA TAT I TTC AAG F N TCC AGG	* GCT CGA A> ACC TGG T> CCT GGA
G G G G G G G G G G G G G G G G G G G	CCA CCA CGT A CCA GCT P	GAA CTT E GCT CGA A	TTG AAC L 102 AAA TTT K ACC TGG	TAT I 20 * CAC GTG H 1070 * AAT TTA N -><	CTC GAG L AAC TTG N GAG CTC E	TTT AAA F  10  TTG AAC L  GCT CGA A	GAG CTC E 	CCC GGG P ATC TAG I CAA GTT OMA	CAA GTT Q 1 TTG AAC L GCG CGC A	GGT CCA G O40 * ACT TGA T 10 ACT TGA T	GGA CCT G AAG TTC K 	CTG GAC L AGG TCC R -Ad c TTC AAG F	* CAA GTT Q 105 TCA AGT S c-1 DC GGGG P	AAC TTG N 50 * AAT TTA N OMAI 100 AAG TTC K	TTC AAG F N	* GCT CGA A> ACC TGG T> CCT GGA

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							25	3/64							
•	1160	· •		11	70 *		1	180		11	190			1200	) *
AAG	GGT	GGA	CAC	TAG	TTG	TAG	TGT	ACG	GAG	TCT	TTA	AGC TCG S	TTC	AGT	CAG
	1:	210			1220	)		123	3Q		124	40 *		12	250
TGT	GAC CTG	GGC CCG	GTT CAA	TAT ATA	GAG CTC	ACC TGG	AGC TCG	TTC AAG	CTC GAG	GTC CAG	AAC TTG	CGT GCA R	GAC CTG	CAT GTA	TCC AGG
		120	<del></del>		1:	270		1	280			129	Ō		
AAG	GTG	AAG TTC	CTG GAC	TCT AGA	TAT ATA	CTC GAG	ACG TGG	TTC AAG	ATC TAG	CCT GGA	TCT AGA	GAT CTA D	GAT CTA	CTG	TAA
1300			1310	)		132	20		133	30		13	340		
TAT ATA	OO 1310  TAT GAC TGC AAG GT ATA CTG ACG TTC CA Y D C K				GAG CTC	CAC GTG	TGG ACC	GGC CCG	CTG GAC	GAG CTC	GAG CTC	CCG GGG	GTT CAA	GAC	TTT
CAC GTG H	TCG ACC N	GAA CTT E	CCT GGA P	GAG CTC E	ATT TAA I	CCA GGT P	GCC CGG A	CCC GGG P	ATG TAC M	TCA AGT S	GAG CTC E	CTG GAC L	ACA TGT T	GAA CTT E	TGA
															)
GTG CAC	GTG CAC	TGT ACA	GCC CGG	CTG GAC	GGG CCC	TTG AAC	TCT AGA	GTG CAC	GGC CCG	CTT GAA	GTG CAC	GGC CCG G	ATC TAG	GTG	GTG CAC
~~~~	1	450			1460	)		14	<del></del>		14	80		14	90
		*	mma	<b>አ</b> ጥረ	Δ T'T'	* CAA				TCA	GGT	* GGC	ACC	TCC	* AGA
GGC CCG G	ACC TGG T	ATC TAG I	AAG	TAG	TAA	GTT	CCG	GAG	GCT	AGT		. CCG G	TGG	AGG S	TC

1500

CAC CCA GGG CCT TTA TGA GTG GGT CCC GGA AAT ACT H P G P L \*> -- I-Ad  $\alpha$ -TM DOMAIN ->



FIG. 29

# FIG. 29A

J. Z	<b>9</b> A	<b>\</b>					01/0	•							
		10			20	)		;	30			<b>40</b>			50 *
CCAC( GGTG(	G TAC	C CGA A	A GA( L	C GTO Q	C TAC I	G GG( P	TC( S	G GAC	GAC L	G GAC	AGT S	Г CGA A	A CGA A	A CAC	C CAC
		6	0 *			70 *			80			9	0		
CAC	GAC	TAC	CAC	GAC	TCG	TCG	GGT	TCC	TGG	TAA	TCA	TAG	AGA	CAG GTC Q	CGA
100							20 *						140		
CAA V	GTG H	CGA A	CGA A	GTG H	CGA A	CTT E	TAG I	TTG N	CTT E	CGA A	CGA G	GCA R	CGA A	AGC TCG S	CCT G>
15	50 *	• • •		160			170							90	
GGG CCC	GGC CCG	CCT	TCG	GGC CCG	GGA CCT	GGG CCC	GGA CCT	AAC TTG	TCC AGG	GAA CTT	AGG TCC	GTA	AAG	GTG CAC V	CAG
	200													24(	
	TTC AAG	TTC	CCG	GAG CTC	TGC ACG	TAC ATG	TAC ATG	ACC TGG	AAC TTG	GGG CCC	ACG TGC	CAG GTC	CGC GCG	ATA TAT I	CGG GCC
	:	250			260			2	70		2	80			
									*						290
			TCT				TTG		CTC					TAC ATG Y	* GAC
GAG	CAC	TGG T	TCT	ATG	TAG I	ATG Y 310	TTG	GCC	GAG CTC	CTC E	ATG	GTG CAC V	GCG R 30	ATG	* GAC CTG
GAG L 	CAC V GAC	TGG T 3	TCT R OO *	ATG Y GAG CTC	TAG I TAC ATG	ATG Y  310 * CGC GCG	TTG N GCG	GCC R GTG CAC	GAG CTC E 320 * ACC TGG	CTC E GAG CTC	ATG Y CTG GAC	GTG CAC V 3:	GCG R 30 *	ATG Y CCA GGT	GAC CTG D>
GAG L AGC TCG	CAC V GAC CTG	TGG T 3 GTG CAC	TCT R OO * GGC CCG	ATG Y GAG CTC E	TAG I TAC ATG	ATG Y 310 * CGC GCG R	TTG N GCG CGC	GCC R GTG CAC	GAG CTC E 320 * ACC TGG T	CTC E GAG CTC	ATG Y CTG GAC	GTG CAC V 3: GGG CCC	GCG R 30 * CGG GCC	ATG Y CCA GGT	GAC CTG

FIG. 29B

29	D					02/0	7-							
0 *		4	00 *			410	)		42	0		4	30	
CAC	CTG D	TGC T	CGC A	ACG C	TCT R	GTG H	TTG N	ATG	CTC	CCC	GGC	CTC	TGG	TCG
440	)					2	160 *			470 *			48,	0
AGG	GAC	GCC	CGG GCC	CTT GAA	GAA CTT	CAG GTC	CCC GGG	AAT TTA	GTC CAG	GCC CGG	ATC TAG	AGG	GAC	AGG
4	190 *			500 *			5	10 *		5	20 *		į	530 *
TGT	CTC	CGG	GAG	TTG	GTG	GTG	TTG	TGA	GAC	CAG	ACA	AGC	CAC	TGT
	5	40		į	550			560	)		57	0		
AAG	ATG	GGT	GCC CGG	AAG TTC	ATC TAG	AAA TTT	GTG CAC	CGC GCG	TGG ACC	TTC AAG	AGG TCC	AAT TTA	CCG	GTC
		590			60	00		6	310 *		(	620 *		
CTC	TGT	CAC	GGG CCC	GTC CAG	TCA AGT	TCC AGG	ACA TGT	CAG GTC	CTT GAA	ATT TAA	TCC	TTA	CCC	CTG
30		(	640 *			650 *			66	60 *		6	70 *	
ACC TGG T	TTC AAG F	GTC	CAG	GAC L	CAG V	TAC M	GAC L	CTC F	TAC M	TGG T	GGA	GTA	CAG GTC Q	CCT
	)		6	90			700			710			72	20
GTC	ATG	TGG	ACG	CAT GTA	CAC	CTC	CAT GTA	GGG	TCG	CTG GAC	TTC	TCG	CCC GGG	ATC TAG
	730			740	)									770
CAC	GAG CTC	ACC	TGA	AGT TCA	GGT CCA	CCG	CCA	CCG	TCG	CCG	CCA	CCA	CCA	TCC AGG
	GTG CAC V  440  * TCC AGG S I-Ad  ACA TGT T  TTC AAG F  GAG CTC E  30  * ACC TGG T  GTGC V  GTGC V  GTGC CAC	* GTG GAC CAC CTG V D  440  * TCC CTG AGG GAC S L I-Ad β-1  490  ACA GAG TGT CTC T E  TTC TAC AAG ATG F Y  GAG ACA CTC TGT E T  30  * ACC TTC TGG AAG T F  680  GTC TAC CAG ATG V Y  730  GTG GAG CAC CTC	GTG GAC ACG CAC CTG TGC V D T  440 * TCC CTG CGG AGG GAC GCC S L R I-Ad β-1 DOM  490  ACA GAG GCC TGT CTC CGG T E A  540  * TTC TAC CCA AAG ATG GGT F Y P  590  GAG ACA GTG CTC TGT CAC E T V  30 * ACC TTC CAG TGG AAG GTC T F Q  680  GTC TAC ACC CAG ATG TGG V Y T  730  GTG GAG TGG CAC CTC ACC	CTG GAC ACG GCG CAC CTG TGC CGC V D T A  440  440  440  440  TCC CTG CGG CGG AGG GAC GCC GCC S L R R I-Ad β-1 DOMAIN  490  ACA GAG GCC CTC TGT CTC CGG GAG T E A L  540  TTC TAC CCA GCC AAG ATG GGT CGG F Y P A  590  GAG ACA GTG GGG CTC TGT CAC CCC E T V G  30  ACC TTC CAG GTC CAG T F Q V  680  ACC TTC CAG GTC CAG T F Q V  680  640  730  640  730  640  730  GTG GAG TGG ACT CAC CTGA  GTG GAG TGG ACG V Y T C  730  GTG GAG TGG ACT CAC CTGA	60	### AUD	### A C	0	### A 10	400	400	A	## A C	## A 10

FIG. 29C

	,. <u> </u>														
		78	30 *		-	790 *			008			81	0 *		
CCA	GGC CCG G	CCG	CCA	AGA S	CCG	CCA G	CCG G	CCA G	AGG S	AGC S	TCA S	CTT E	CTG	CTG	TAA
820			830			84	40 *		8	350 *			860		
GAG CTC	GCC CGG A I-A	CTG	CAC	GTA CAT	GGC CCG	TTC AAG	TAT ATA	GGT CCA	ACA TGT	ACT TGA	GTT CAA	TAT ATA	CAG GTC	AGA	CCT GGA P>
8	70 *		8	80			890	1		90	0		9	10	
GGA CCT G	GAC CTG	TAA	GGC	CAG GTC	TAC ATG	ACA TGT	CAT GTA	GAA CTT	TTT AAA	GAT CTA	GGT CCA	GAT CTA	GAG CTC	TTG AAC	AAG
<b></b> -	920			9:	30			940			950			96	§0
ATA	GTG CAC V	CTG	AAC	CTA	TTC	TTC	TTT	ACT TGA	CAG	ACC	TCC	GAA	GGA	CTC	AAA
	ç	970			980			99	90		10	000			1010
CCG	CAA GTT Q	AAC	ATA TAT	CTC GAG	TTT AAA	GAG CTC	CCC GGG	CAA GTT	GGT CCA	GSA CCT	CTG GAC	CAA GTT	AAC TTG	ATA TAT	GCT CGA
		10	20		1	030			1040	)		10	50 *		
CGT	GAA CTT E	AAA TTT	CAC GTG	AAC TTG	TTG AAC	GGA CCT	ATC TAG	TTG AAC	ACT TGA	AAG TTC	AGG TCC	TCA AGT	AAT TTA	AAG	TGG
1060	)		1070	)		10	080		1	090		1	1100		
CCA	GCT CGA A	ACC	AAT	GAG	GCT	CCT	CAA	GCG	ACT	GTG	TTC	CCC	AAG	TCC	CCT GGA P>
	110														
GTG	CTG	CTG	GGT	CAG	CCC	AAC	ACC	CTT	ATC	TGC	TTT	GTG	GAC	AAC	ATC TAG I>

11 <u>6</u> 0				1	170 *		1180 *			1190 *				1200	
AAG	GGT	CCT GGA P	CAC	TAG	TTG	TAG	TGT	ACS	GAG	TCT	TTA	TCG	TTC	AGT	CAG
12 <u>1</u> 0				1220				1230			1240				1250
TGT	CTG	GGC CCG G	CAA	ATA	CTC	TGG	TCG	AAG	GAG	CAG	TTG	GCA	CTG	GTA	AGG
	1260				1270 *			1280 *			1290 *				
AAG	GTG	AAG TTC K	GAC	AGA	ATA	GAG	TGG	AAG	TAG	GGA	AGA	CTA	CTA	CTG	TAA
1300	300 1310				1320 *				1330			1340			
TAT ATA	GAC CTG	TGC ACG C	AAG TTC K	GTG CAC V	GAG CTC E	CAC GTG H	TGG ACC W	GGC CSS G	CTG GAC L	GAG CTC E	GAG CTC E	CCG GGC P	GTT CAA V	CTG GAC L	TTT K>
I-Ad $\alpha$ -2 DOMAIN															
13	1360 *			1370			)	138			80 *				
CAC GTG H	TGG ACC W	GAG CTC E ><	GAA CTT E	GAA CTT E	GAG CTC E	TAC ATG Y	ATG TAC M	CCG GGC P	ATG TAC M	GAA CTT E	TGA ACT *>				

FIG. 29D